

(19) World Intellectual Property Organization
International Bureau



(43) International Publication Date
22 February 2001 (22.02.2001)

PCT

(10) International Publication Number
WO 01/11966 A1

(51) International Patent Classification⁷: **A01N 43/40**,
43/82, 43/80, 43/54, 43/56, 43/90, 43/50, 43/78, 43/42,
43/60, 53/00, 47/38, 47/40, 47/24

Cambridge CB1 6LW (GB). **TURNER, Richar, Michäel**
[GB/GB]: 3 Harebell Close, Cherry Hinton, Cambridge
CB1 4YL (GB).

(21) International Application Number: PCT/EP00/08269

(74) Agent: **MERIGEAULT, Shona**; Aventis CropScience
S.A., Département Propriété Industrielle, B.P. 9163,
F-69263 Lyon Cedex 09 (FR).

(22) International Filing Date: 11 August 2000 (11.08.2000)

(25) Filing Language: English

(81) Designated States (*national*): AE, AG, AL, AM, AT, AU,
AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ,
DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR,
HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ,
NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM,
TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW.

(26) Publication Language: English

(30) Priority Data:
9919588.5 18 August 1999 (18.08.1999) GB

(71) Applicant (*for all designated States except US*): **AVEN-
TIS CROPSOURCE GMBH** [DE/DE]: Brüningstrasse
50, D-65929 Frankfurt am Main (DE).

(84) Designated States (*regional*): ARIPO patent (GH, GM,
KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW). Eurasian
patent (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM). European
patent (AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE,
IT, LU, MC, NL, PT, SE). OAPI patent (BF, BJ, CF, CG,
CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG).

(72) Inventors; and

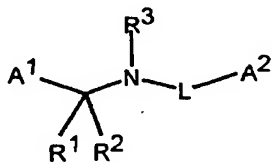
(75) Inventors/Applicants (*for US only*): **COOKE, Tracey**
[GB/GB]; 7 Larch Avenue, Brickett Wood, St. Albans
AL2 3SN (GB). **EKWURU, Tennyson** [GB/FR]; Aventis
CropScience SA, 14-20, rue Pierre Baizet, F-69263 Lyon
Cedex 09 (FR). **HARDY, David** [GB/GB]; 46 St. Bedes
Gardens, Cambridge CB1 3UF (GB). **MILLWARD, Peter**
[GB/GB]; 28 Holly Trees, Bar Hill, Cambridge CB3 8SF
(GB). **MOLONEY, Brian** [GB/GB]; 2 Crookdale Beck,
Oxon OX11 7US (GB). **PETTINGER, Andrew** [GB/FR];
17, rue Saint Lazare, F-69007 Lyon (FR). **THOMAS,
Peter, Stanley** [GB/GB]; 94 Balsham Road, Linton,

Published:

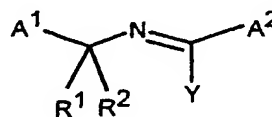
- With international search report.
- Before the expiration of the time limit for amending the
claims and to be republished in the event of receipt of
amendments.

For two-letter codes and other abbreviations, refer to the "Guid-
ance Notes on Codes and Abbreviations" appearing at the begin-
ning of each regular issue of the PCT Gazette.

(54) Title: FUNGICIDES



(I)



(II)

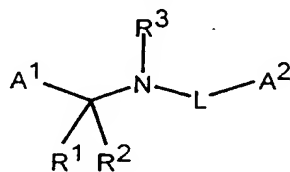
(57) Abstract: Use of compounds of general formula (I) or (II) or salts thereof as phytopathogenic fungicides wherein the various radicals and substituents are as defined in the description, pesticidal compositions containing them and method for combating pests which comprises applying these.

WO 01/11966 A1

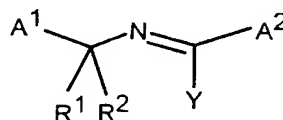
Fungicides

This invention relates to compounds having fungicidal activity.

- 5 In a first aspect the invention provides the use of compounds of general formula I or II or salts thereof as phytopathogenic fungicides



(I)



(II)

wherein

- 10 A¹ is 2-pyridyl or its *N*-oxide, each of which may be substituted by up to four groups at least one of which is haloalkyl;

A² is heterocyclyl or carbocyclyl, each of which may be substituted (A² is preferably optionally substituted heterocyclyl or optionally substituted phenyl);

- 15 R¹ and R², which may be the same or different, are R^b, cyano, nitro, halogen, -OR^b, -SR^b or optionally substituted amino, or R¹ and R² together with the carbon to which they are attached may form a 3-, 4-, 5- or 6- carbo- or heterocyclic ring, which may be substituted (R¹ and R² are preferably hydrogen, acyl, optionally substituted alkyl or cyano);

- 20 R³ is R^b, -OR^b, or -N(R^b)₂, cyano, *N*-substituted iminomethyl or nitro; or R³ and A², together with the interconnecting atoms, may form a 5- or 6-membered ring (R³ is preferably hydrogen, *N*-substituted iminomethyl or optionally substituted alkyl);

L is -C(=X)- or -SO₂-, where X is oxygen, sulfur, N-OR^b, N-R^b or N-N(R^b)₂ (L is preferably -C(=O)-, -C(=S)- or -C(=NOR^b)-); and

- 25 Y is halogen, -OR^b, -SR^b, -N(R^b)₂, -NR^b(OR^b) or -NR^bN(R^b)₂ (preferably -OR^b, -SR^b or -N(R^b)₂);

and R^b, which may be the same or different, is alkyl, alkenyl, alkynyl, carbocyclyl or heterocyclyl, each of which may be substituted: or hydrogen or acyl, or two

adjacent R^b groups together with the interconnecting atoms, may form a 5- or 6-membered ring;

with the proviso that when A¹ is 2-pyridyl, R¹ is hydrogen. R² is hydrogen, optionally substituted alkyl or acyl. L is -C(=X)- or -SO₂-, X is oxygen or sulfur and R³ is hydrogen or optionally substituted alkyl, A² is not optionally substituted phenyl.

Preferred substituents on the 2-pyridyl group (A¹) are halogen, hydroxy, cyano, nitro, SF₅, trialkylsilyl, optionally substituted amino, acyl, or a group -R^a, -OR^a or -SR^a, or a group -C(R^a)=N-Q, where Q is -R^a, -OR^a, -SR^a or optionally substituted amino, wherein R^a is alkyl, alkenyl, alkynyl, carbocyclyl or heterocyclyl, each of which may be substituted; or two adjacent substituents together with the atoms to which they are attached form an optionally substituted ring which can contain up to 3 hetero atoms. Especially preferred substituents are alkoxy, alkyl, cyano, halogen, nitro, alkoxycarbonyl, alkylsulfinyl, alkylsulfonyl and trifluoromethyl, particularly chlorine and trifluoromethyl.

Preferably, the 2-pyridyl group is substituted at the 3 and/or 5 position.

The invention also includes any of the compounds specifically exemplified hereinafter.

Any alkyl group may be straight or branched and is preferably of 1 to 10 carbon atoms, especially 1 to 7 and particularly 1 to 5 carbon atoms.

Any alkenyl or alkynyl group may be straight or branched and is preferably of 2 to 7 carbon atoms and may contain up to 3 double or triple bonds which may be conjugated, for example vinyl, allyl, butadienyl or propargyl.

Any carbocyclyl group may be saturated, unsaturated or aromatic, and contain 3 to 8 ring-atoms. Preferred saturated carbocyclyl groups are cyclopropyl, cyclopentyl or cyclohexyl. Preferred unsaturated carbocyclyl groups contain up to 3 double bonds. A preferred aromatic carbocyclyl group is phenyl. The term carbocyclic should be similarly construed. In addition, the term carbocyclyl includes any fused combination of carbocyclyl groups, for example naphthyl, phenanthryl, indanyl and indenyl.

Any heterocyclyl group may be saturated, unsaturated or aromatic, and contain 5 to 7 ring-atoms up to 4 of which may be hetero-atoms such as nitrogen, oxygen and sulfur. Examples of heterocyclyl groups are furyl, thienyl, pyrrolyl, pyrrolinyl, pyrrolidinyl, imidazolyl, dioxolanyl, oxazolyl, thiazolyl, imidazolyl, imidazoliny, imidazolidinyl, pyrazolyl, pyrazolinyl, pyrazolidinyl, isoxazolyl, isothiazolyl, oxadiazolyl, triazolyl, thiadiazolyl, pyranyl, pyridyl, piperidinyl, dioxanyl, morpholino, dithianyl, thiomorpholino, pyridazinyl, pyrimidinyl, pyrazinyl, piperazinyl, sulfolanyl, tetrazolyl, triazinyl, azepinyl, oxazepinyl, thiazepinyl, diazepinyl and thiazolinyl. In addition, the term heterocyclyl includes fused heterocyclyl groups, for example benzimidazolyl, benzoxazolyl, imidazopyridinyl, benzoxazinyl, benzothiazinyl, oxazolopyridinyl, benzofuranyl, quinolinyl, quinazolinyl, quinoxalinyl, dihydroquinazolinyl, benzothiazolyl, phthalimido, benzofuranyl, benzodiazepinyl, indolyl and isoindolyl. The term heterocyclic should be similarly construed.

Any alkyl, alkenyl, alkynyl, carbocyclyl or heterocyclyl group, when substituted, may be substituted by one or more substituents, which may be the same or different, and may be selected from the list: hydroxy; mercapto; azido; nitro; halogen; cyano; acyl; optionally substituted amino; optionally substituted carbocyclyl; optionally substituted heterocyclyl; cyanato; thiocyanato; $-SF_5$; $-OR^a$; $-SR^a$ and $-Si(R^a)_3$, where R^a is alkyl, alkenyl, alkynyl, carbocyclyl or heterocyclyl, each of which may be substituted. In the case of any carbocyclyl or heterocyclyl group the list includes additionally: alkyl, alkenyl and alkynyl, each of which may be substituted. Preferred substituents on any alkyl, alkenyl or alkynyl group are alkoxy, haloalkoxy or alkylthio, each containing 1 to 5 carbon atoms; halogen; or optionally substituted phenyl. Preferred substituents on any carbocyclyl or heterocyclyl group are alkyl, haloalkyl, alkoxy, haloalkoxy or alkylthio, each containing 1 to 5 carbon atoms; halogen; or optionally substituted phenyl.

In the case of any alkyl group or any unsaturated ring-carbon in any carbocyclyl or heterocyclyl group the list includes a divalent group such as oxo or imino, which may be substituted by optionally substituted amino, R^a or $-OR^a$. Preferred groups are oxo, imino, alkylimino, oximino, alkylloximino or hydrazono.

Any amino group, when substituted and where appropriate, may be substituted by one or two substituents which may be the same or different, selected from the list: optionally substituted alkyl, optionally substituted amino, $-OR^a$ and acyl groups. Alternatively two

substituents together with the nitrogen to which they are attached may form a heterocyclyl group, preferably a 5 to 7-membered heterocyclyl group, which may be substituted and may contain other hetero atoms, for example morpholino, thiomorpholino or piperidinyl.

5

The term acyl includes the residues of sulfur and phosphorus-containing acids as well as carboxylic acids. Typically the residues are covered by the general formulae $-C(=X^a)R^c$, $-S(O)_pR^c$ and $-P(=X^a)(OR^a)(OR^a)$, where appropriate X^a is O or S. R^c is as defined for R^a , $-OR^a$, $-SR^a$, optionally substituted amino or acyl; and p is 1 or 2. Preferred groups are $-C(=O)R^d$, $-C(=S)R^d$, and $-S(O)_pR^d$ where R^d is alkyl, C_1 to C_5 alkoxy, C_1 to C_5 alkylthio, phenyl, heterocyclyl or amino, each of which may be substituted.

10

Complexes of compounds of the invention are usually formed from a salt of formula MAN_2 , in which M is a divalent metal cation, e.g. copper, manganese, cobalt, nickel, iron or zinc and An is an anion, e.g. chloride, nitrate or sulfate.

15

In cases where the compounds of the invention exist as the E and Z isomers, the invention includes individual isomers as well as mixtures thereof.

20

In cases where compounds of the invention exist as tautomeric isomers, the invention includes individual tautomers as well as mixtures thereof.

In cases where the compounds of the invention exist as optical isomers, the invention includes individual isomers as well as mixtures thereof.

25

The compounds of the invention have activity as fungicides, especially against fungal diseases of plants, e.g. mildews and particularly cereal powdery mildew (*Erysiphe graminis*) and vine downy mildew (*Plasmopara viticola*), rice blast (*Piricularia oryzae*), cereal eyespot (*Pseudocercospora herpotrichoides*), rice sheath blight (*Pellicularia sasakii*), grey mould (*Botrytis cinerea*), damping off (*Rhizoctonia solani*), wheat brown rust (*Puccinia recondita*), late tomato or potato blight (*Phytophthora infestans*), apple scab (*Venturia inaequalis*), and glume blotch (*Leptosphaeria nodorum*). Other fungi against which the compounds may be active include other powdery mildews, other rusts, and other general pathogens of Deuteromycete, Ascomycete, Phycomycete and

30

Basidiomycete origin.

5 The invention thus also provides a method of combating fungi at a locus infested or liable to be infested therewith, which comprises applying to the locus a compound of formula I.

The invention also provides an agricultural composition comprising a compound of formula I in admixture with an agriculturally acceptable diluent or carrier.

10 The composition of the invention may of course include more than one compound of the invention.

15 In addition, the composition can comprise one or more additional active ingredients, for example compounds known to possess plant-growth regulant, herbicidal, fungicidal, insecticidal, acaricidal, antimicrobial or antibacterial properties. Alternatively the compound of the invention can be used in sequence with the other active ingredient.

20 The diluent or carrier in the composition of the invention can be a solid or a liquid optionally in association with a surface-active agent, for example a dispersing agent, emulsifying agent or wetting agent. Suitable surface-active agents include anionic compounds such as a carboxylate, for example a metal carboxylate of a long chain fatty acid; an *N*-acylsarcosinate; mono- or di-esters of phosphoric acid with fatty alcohol ethoxylates or alkyl phenol ethoxylates or salts of such esters; fatty alcohol sulfates such as sodium dodecyl sulfate, sodium octadecyl sulfate or sodium cetyl sulfate; ethoxylated fatty alcohol sulfates; ethoxylated alkylphenol sulfates; lignin sulfonates; petroleum sulfonates; alkyl-aryl sulfonates such as alkyl-benzene sulfonates or lower alkyl naphthalene sulfonates, e.g. butyl-naphthalene sulfonate; salts of sulfonated naphthalene-formaldehyde condensates; salts of sulfonated phenol-formaldehyde condensates; or more complex sulfonates such as the amide sulfonates, e.g. the sulfonated condensation product of oleic acid and *N*-methyl taurine; the dialkyl sulfosuccinates, e.g. the sodium sulfonate of dioctyl succinate; acid derivatives of alkyl glycosides and alkylpolyglycosides materials and their metal salts, e.g. alkyl polyglycoside citrate or tartrate materials; or mono-, di- and tri-alkyl esters of citric acid and their metal salts.

35

Nonionic agents include condensation products of fatty acid esters, fatty alcohols, fatty acid amides or fatty-alkyl- or alkenyl-substituted phenols with ethylene and/or propylene oxide; fatty esters of polyhydric alcohol ethers, e.g. sorbitan fatty acid esters; condensation products of such esters with ethylene oxide, e.g. polyoxyethylene sorbitan fatty acid esters; alkyl glycosides, alkyl polyglycoside materials; block copolymers of ethylene oxide and propylene oxide; acetylenic glycols such as 2,4,7,9-tetramethyl-5-decyne-4,7-diol, ethoxylated acetylenic glycols; acrylic based graft copolymers; alkoxylated siloxane surfactants; or imidazoline type surfactants, e.g. 1-hydroxyethyl-2-alkylimidazoline.

Examples of a cationic surface-active agent include, for instance, an aliphatic mono-, di-, or polyamine as an acetate, naphthenate or oleate; an oxygen-containing amine such as an amine oxide, polyoxyethylene alkylamine or polyoxypropylene alkylamine; an amide-linked amine prepared by the condensation of a carboxylic acid with a di- or polyamine; or a quaternary ammonium salt.

The compositions of the invention can take any form known in the art for the formulation of agrochemicals, for example, a solution, an aerosol, a dispersion, an aqueous emulsion, a microemulsion, a dispersible concentrate, a dusting powder, a seed dressing, a fumigant, a smoke, a dispersible powder, an emulsifiable concentrate, granules or an impregnated strip. Moreover it can be in a suitable form for direct application or as a concentrate or primary composition which requires dilution with a suitable quantity of water or other diluent before application.

A dispersible concentrate comprises a compound of the invention dissolved in one or more water miscible or semi-water miscible solvents together with one or more surface active and/or polymeric material. Addition of the formulation to water results in the crystallisation of the active ingredient, the process being controlled by the surfactants and/or polymers resulting in a fine dispersion.

A dusting powder comprises a compound of the invention intimately mixed and ground with a solid pulverulent diluent, for example, kaolin.

An emulsifiable concentrate comprises a compound of the invention dissolved in a water-immiscible solvent which forms an emulsion or microemulsion on addition to water in the presence of an emulsifying agent.

5 A granular solid comprises a compound of the invention associated with similar diluents to those that may be employed in dusting powders, but the mixture is granulated by known methods. Alternatively it comprises the active ingredient absorbed or coated on a pre-formed granular carrier, for example. Fuller's earth, attapulgite, silica or limestone grit.

10

Wettable powders, granules or grains usually comprise the active ingredient in admixture with suitable surfactants and an inert powder diluent such as clay or diatomaceous earth.

15 Another suitable concentrate is a flowable suspension concentrate which is formed by grinding the compound with water or other liquid, surfactants and a suspending agent.

The concentration of the active ingredient in the composition of the present invention, as applied to plants is preferably within the range of 0.0001 to 1.0 per cent by weight, especially 0.0001 to 0.01 per cent by weight. In a primary composition, the amount of
20 active ingredient can vary widely and can be, for example, from 5 to 95 per cent by weight of the composition.

The invention is generally applied to seeds, plants or their habitat. Thus, the compound can be applied directly to the soil before, at or after drilling so that the presence of active
25 compound in the soil can control the growth of fungi which may attack seeds. When the soil is treated directly the active compound can be applied in any manner which allows it to be intimately mixed with the soil such as by spraying, by broadcasting a solid form of granules, or by applying the active ingredient at the same time as drilling by inserting it in the same drill as the seeds. A suitable application rate is within the range of from 5 to
30 1000 g per hectare, more preferably from 10 to 500 g per hectare.

Alternatively the active compound can be applied directly to the plant by, for example, spraying or dusting either at the time when the fungus has begun to appear on the plant or before the appearance of fungus as a protective measure. In both such cases the
35 preferred mode of application is by foliar spraying. It is generally important to obtain

good control of fungi in the early stages of plant growth, as this is the time when the plant can be most severely damaged. The spray or dust can conveniently contain a pre- or post-emergence herbicide if this is thought necessary. Sometimes, it is practicable to treat the roots, bulbs, tubers or other vegetative propagule of a plant before or during planting, for example, by dipping the roots in a suitable liquid or solid composition. When the active compound is applied directly to the plant a suitable rate of application is from 0.025 to 5 kg per hectare, preferably from 0.05 to 1 kg per hectare.

In addition, the compounds of the invention can be applied to harvested fruits, vegetables or seeds to prevent infection during storage.

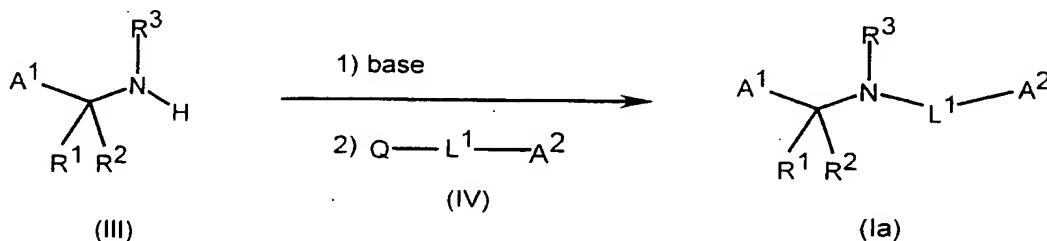
In addition, the compounds of the invention can be applied to plants or parts thereof which have been genetically modified to exhibit a trait such as fungal and/or herbicidal resistance.

In addition the compounds of the invention can be used to treat fungal infestations in timber and in public health applications. Also the compounds of the invention can be used to treat insect and fungus infestations in domestic and farm animals.

Compounds of the invention may be prepared, in known manner, in a variety of ways.

Compounds of formula Ia, i.e. compounds of general formula I where L is L¹ which is -C(=O)-, -C(=S)-, -SO₂- or -C(=NOH)- may be prepared according to reaction scheme 1 by reacting compounds of formula III or their hydrochloride salt with compounds of formula IV, where Q is a leaving group such as halogen, preferably chlorine. A preferred base is triethylamine.

Scheme 1



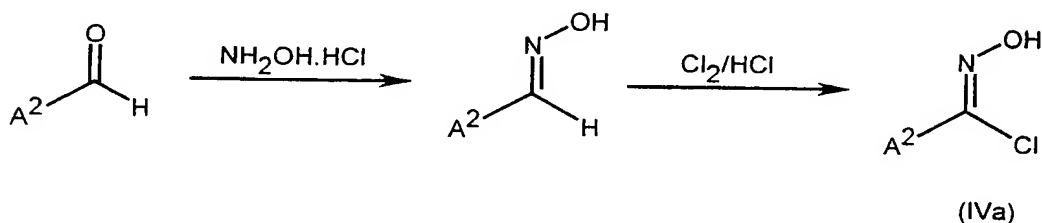
Compounds of formula IV where L¹ is -C(=O)-, -C(=S)- or -SO₂- can be prepared from the corresponding hydroxy compound by methods known to the skilled chemist.

Compounds of formula IV can be isolated and used according to scheme 1. Alternatively, IV may be generated *in situ* by methods known to the skilled chemist, for example, using POCl₃ to generate the acid chloride from the corresponding carboxylic acid, followed by addition of III.

5

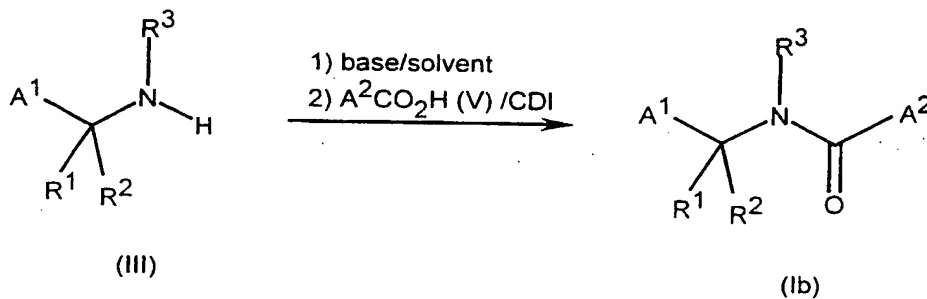
Compounds of formula IVa, i.e. compounds of general formula IV where L¹ is –C(=NOH)– can be prepared according to reaction scheme 2.

Scheme 2



Compounds of formula Ib, i.e. compounds of general formula I where L is –C(=O)– may be prepared according to reaction scheme 3 by reacting compounds of formula III in the presence of a suitable base such as triethylamine with compounds of formula V in the presence of carbonyl diimidazole (CDI).

Scheme 3



15

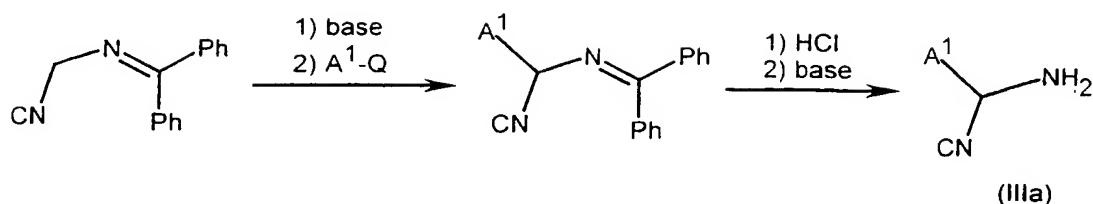
The preparation of compounds of formula V where A² is 3-hydroxy-2-benzo[b]furyl form part of the state of the art see P C Unangst, D T Connor, S R Miller, *J. Het. Chem.* 1996, 33, 2025-2030.

20

Many compounds of formula III may be prepared by methods described in international application PCT/GB/99/00304. Compounds of formula IIIa, i.e. compounds of general formula III where R¹ is hydrogen and R² is cyano, may be prepared by methods analogous to those described therein (see reaction scheme 3a).

10

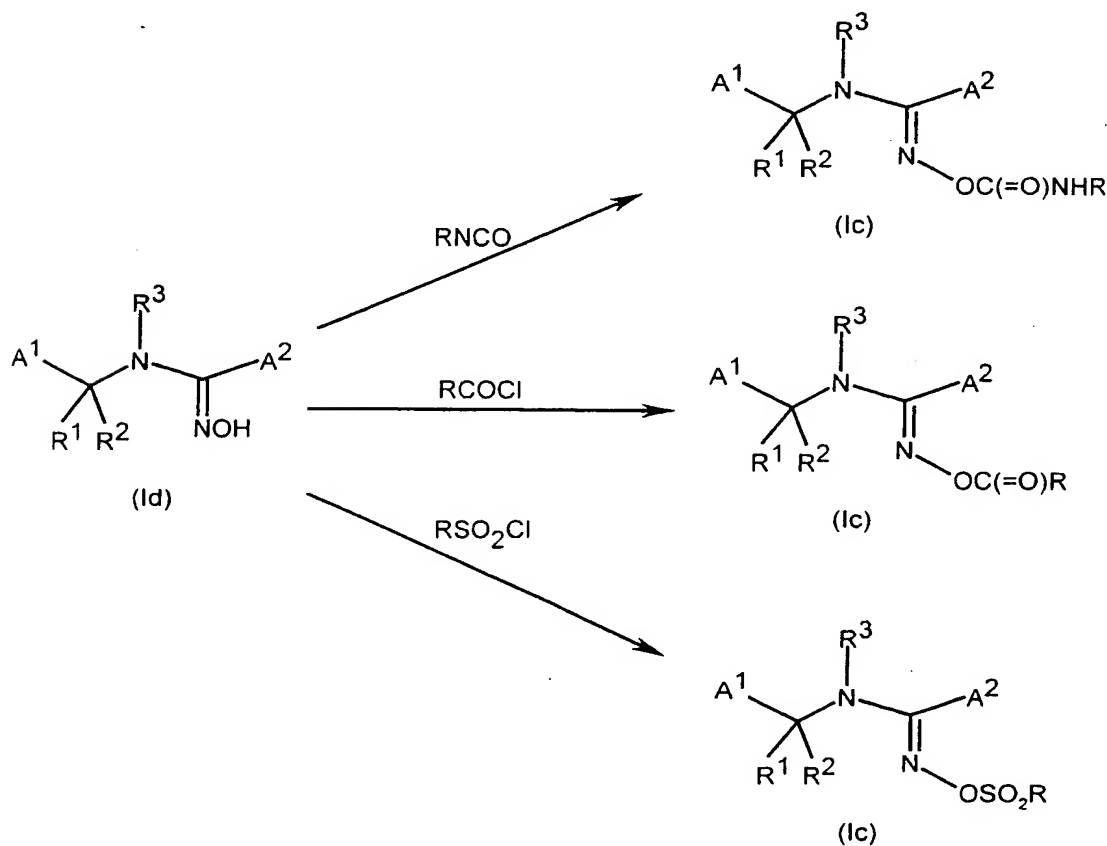
Scheme 3a



Compounds of formula III where R^1 is alkyl and R^2 is cyano or acyl, may be prepared by alkylating analogues where R^1 is hydrogen.

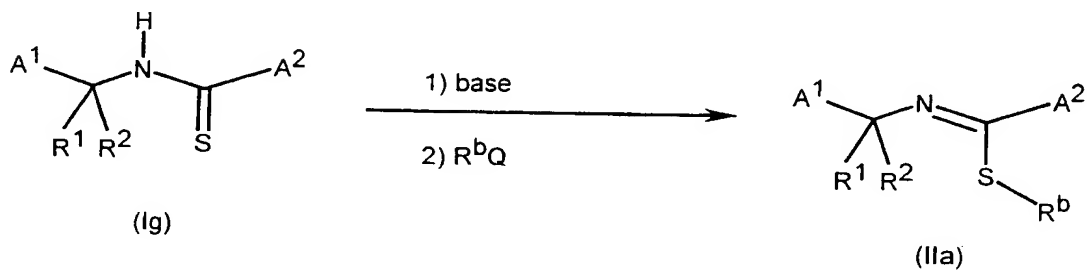
Compounds of formula Ic, i.e. compounds of general formula I where L is $-C(=N-OR^b)-$ may be prepared from compounds of formula Id where L^1 is $-C(=NOH)-$ according to Scheme 4 using methodology known to the skilled chemist. For example compounds of formula Ic where R^b is $-C(=O)NHR$ may be prepared by reaction with $R-NCO$; compounds where R^b is $-C(=O)R$ may be prepared by reaction with $RCOCl$ and compounds where R^b is $-SO_2R$ may be prepared by reaction with RSO_2Cl .

Scheme 4

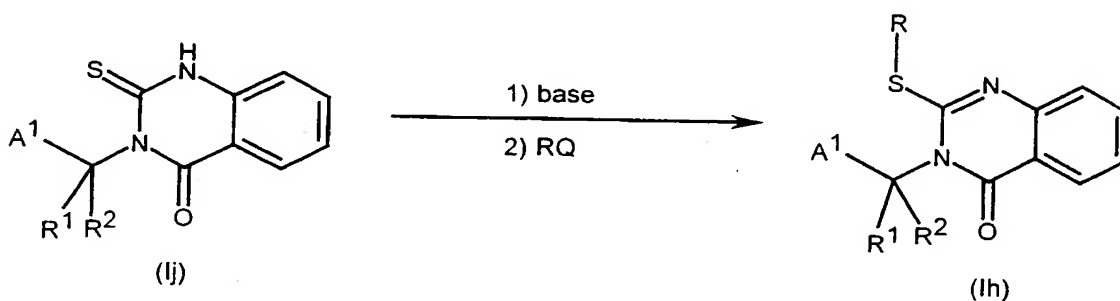


15

Compounds of formula IIa, i.e. compounds of general formula II where Y is $-SR^b$ may be prepared from compounds of formula Ig, i.e. compounds of formula Ia where L^1 is $-C(=S)-$ according to reaction scheme 5. Reaction conditions comprise treating Ig with a base such as sodium hydride followed by reaction with R^bQ where Q is a leaving group preferably halogen.

Scheme 5

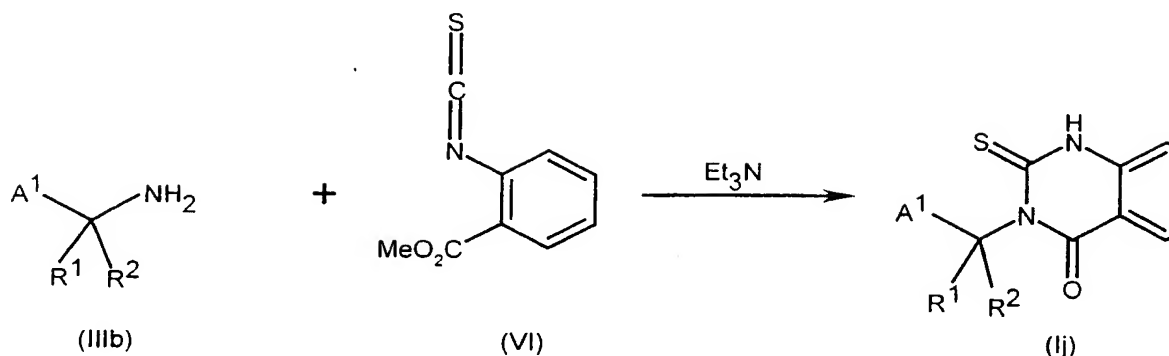
Compounds of formula Ih, i.e. compounds of general formula I where A^2 is 2-substituted phenyl which substituent together with R^3 and the interconnecting atoms forms a 6-membered ring, may be prepared from compounds of formula Ij by treatment with base, preferably potassium carbonate in acetone, followed by RQ where Q is a leaving group, according to reaction scheme 6.

Scheme 6

Compounds of general formula Ij, may be prepared by reacting compounds of formula IIIb with compounds of formula VI in the presence of a suitable base, such as triethylamine, according to reaction scheme 7.

12

Scheme 7



Compounds of formula VI may be prepared by methods known to the skilled chemist from the corresponding amino compound.

Compounds of formula I or II where A¹ is pyridyl N-oxide, may be prepared from the corresponding pyridyl derivative by reactions known to the skilled chemist; for example reaction with peracetic acid.

Other methods will be apparent to the chemist skilled in the art, as will be the methods for preparing starting materials and intermediates.

Collections of compounds of formula I and II may also be prepared in a parallel manner, either manually, automatically or semi-automatically. This parallel preparation may be applied to the reaction procedure, work-up or purification of products or intermediates. For a review of such procedures see by S.H. DeWitt in "Annual Reports in Combinatorial Chemistry and Molecular Diversity: Automated synthesis", Volume 1, Verlag Escom 1997, pages 69 to 77.

Furthermore, compounds of the formula I or II may be prepared using solid-supported methods, where the reactants are bound to a synthetic resin. See for example: Barry A. Bunin in "The Combinatorial Index", Academic Press, 1998 and "The tea-bag method" (Houghten, US 4,631,211; Houghten et al., Proc. Natl. Acad. Sci, 1985, 82, 5131-5135).

The invention is illustrated in the following Examples. Structures of isolated, novel compounds were confirmed by NMR and/or other appropriate analyses.

Example 1

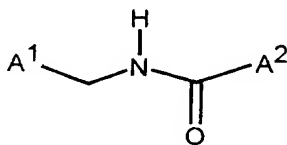
N-[(3-Chloro-5-trifluoromethyl-2-pyridyl)methyl]-2-furamide(Compound 1)

To a mixture of (3-chloro-5-trifluoromethyl-2-pyridyl)methylamine (1 mmol) in tetrahydrofuran (5 ml) was added triethylamine (1 mmol) at room temperature and the mixture was stirred at room temperature for 1.5 hour. The mixture was added to a solution of 2-furoyl chloride (1 mmol) in tetrahydrofuran (5 ml) at room temperature and left to stir at room temperature overnight. The mixture was evaporated to dryness and the residue washed with water. The solid was filtered, washed with diethyl ether/ light petroleum (b.p. 60-80 °C) and dried to give the title product. ¹H N.M.R. (CDCl₃) δ(ppm) 4.9 (2H, d), 6.55 (1H, m), 7.2 (1H, m), 7.5 (1H, s), 7.8 (1H, br.s), 8.0 (1H, s) and 8.8 (1H, s).

Example 2N-[(3-Chloro-5-trifluoromethyl-2-pyridyl)methyl]-5-bromofuramide(Compound 23)

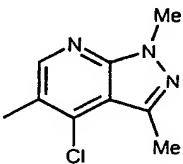
To a mixture of 5-bromofuroic acid (0.19 g) and carbonyl diimidazole (CDI) in dichloromethane was added (3-chloro-5-trifluoromethyl-2-pyridyl)methylamine and the mixture was stirred at room temperature overnight. The mixture was washed with 2M hydrochloric acid, then saturated sodium bicarbonate, dried (MgSO₄) and evaporated to give the title product, m.p. 77.8 °C.

The following compounds of formula Ik (see Table A), i.e. compounds of general formula I where L is -C(=O)- and R¹, R² and R³ are hydrogen, may be prepared by methods analogous to those of Examples 1 and 2.



(Ik)

Table A

Cmp	A ¹	A ²	m.p.(°C)
1	3-Cl-5-CF ₃ -2-pyridyl	2-furyl	oil
2	3-Cl-5-CF ₃ -2-pyridyl	2-thienyl	oil
3	3-Cl-5-CF ₃ -2-pyridyl	3,5-diMe-isoxazol-4-yl	oil
4	3-Cl-5-CF ₃ -2-pyridyl	5-Me-1,2,3-thiadiazol-4-yl	oil
5	3-Cl-5-CF ₃ -2-pyridyl	6-Cl-3-pyridyl	oil
6	3-Cl-5-CF ₃ -2-pyridyl	5-Cl-6-MeO-3-pyridyl	oil
7	3-Cl-5-CF ₃ -2-pyridyl	2-Cl-4-CF ₃ -pyrimidin-5-yl	oil
8	3-Cl-5-CF ₃ -2-pyridyl	1-Ph-5-CF ₃ -pyrazol-4-yl	oil
9	3-Cl-5-CF ₃ -2-pyridyl	2-Cl-3-pyridyl	oil
10	3-Cl-5-CF ₃ -2-pyridyl	3-pyridyl	oil
11	3-Cl-5-CF ₃ -2-pyridyl	3-Cl-2-thienyl	oil
12	3-Cl-5-CF ₃ -2-pyridyl	2-quinazoline	oil
13	3-Cl-5-CF ₃ -2-pyridyl	3-Cl-2-benzo[b]thienyl	oil
14	3-Cl-5-CF ₃ -2-pyridyl		oil
15	3-Cl-5-CF ₃ -2-pyridyl	2-PhO-3-pyridyl	oil
16	3-Cl-5-CF ₃ -2-pyridyl	2-Me-5-(4-Cl-phenyl)-3-furyl	oil
17	3-Cl-5-CF ₃ -2-pyridyl	5,6-diCl-3-pyridyl	171-3
18	3-Cl-5-CF ₃ -2-pyridyl	1-(4-F-2-CF ₃ -benzyl)-imidazol-2-yl	121
19	3-Cl-5-CF ₃ -2-pyridyl	2-pyridyl	112
20	3-Cl-5-CF ₃ -2-pyridyl	5-(3,5-diCl-phenoxy)-2-furyl	107
21	3-Cl-5-CF ₃ -2-pyridyl	4,5-DiBr-2-thienyl	125
22	3-Cl-5-CF ₃ -2-pyridyl	isoxazol-5-yl	110-1
23	3-Cl-5-CF ₃ -2-pyridyl	5-Br-2-furyl	77-8
24	3-Cl-5-CF ₃ -2-pyridyl	3-Me-2-thienyl	134-5
25	3-Cl-5-CF ₃ -2-pyridyl	5-MeO-2-benzo[b]thienyl	122-3
26	3-Cl-5-CF ₃ -2-pyridyl	5-NO ₂ -2-benzo[b]thienyl	149-51
27	3-Cl-5-CF ₃ -2-pyridyl	5-Me-2-thienyl	129-30

Cmp	A ¹	A ²	m.p.(°C)
28	3-Cl-5-CF ₃ -2-pyridyl	2,4-diMe-5-thiazolyl	103-4
29	3-Cl-5-CF ₃ -2-pyridyl	1-diMe-sulfamoylimidazol-4-yl	159-60
30	3-Cl-5-CF ₃ -2-pyridyl	3,5-diMe-1-Ph-imidazol-4-yl	158-9
31	3-Cl-5-CF ₃ -2-pyridyl	1-Ph-imidazol-4-yl	126-7
32	3-Cl-5-CF ₃ -2-pyridyl	4,6-diMeO-2-(4-Cl- — , — -diMe-benzyl)pyrimidin-5-yl	oil
33	3-Cl-5-CF ₃ -2-pyridyl	5-Br-3-pyridyl	131-3
34	3-Cl-5-CF ₃ -2-pyridyl	2-MeS-3-pyridyl	131-2
35	3-Cl-5-CF ₃ -2-pyridyl	2-MeO-3-pyridyl	149-50
36	3-Cl-5-CF ₃ -2-pyridyl	6-MeO-3-pyridyl	114-5
37	3-Cl-5-CF ₃ -2-pyridyl	2-Cl-6-Me-3-pyridyl	108-9
38	3-Cl-5-CF ₃ -2-pyridyl	2-phenylthiomethylthio-3-pyridyl	110-1
39	3-Cl-5-CF ₃ -2-pyridyl	2-Cl-4-pyridyl	109-10
40	3-Cl-5-CF ₃ -2-pyridyl	2-Ph-quinolin-4-yl	152-3
41	3-Cl-5-CF ₃ -2-pyridyl	2,6-diMeO-3-pyridyl	181
42	3-Cl-5-CF ₃ -2-pyridyl	1-Me-3-indolyl	171
43	3-Cl-5-CF ₃ -2-pyridyl	3-2H-benzopyranyl	123
44	3-Cl-5-CF ₃ -2-pyridyl	4,6-diMeO-pyrimidin-2-yl	200
45	3-Cl-5-CF ₃ -2-pyridyl	4-CF ₃ -3-pyridyl	oil
46	3-Cl-5-CF ₃ -2-pyridyl	1-Me-4,5-diBr-2-pyrollyl	oil
47	3-Cl-5-CF ₃ -2-pyridyl	4,5-diBr-2-pyrollyl	oil
48	3-Cl-5-CF ₃ -2-pyridyl	4-pyridyl	124-5
49	3-Cl-5-CF ₃ -2-pyridyl	5-Me-pyrazin-2-yl	118-9
50	3-Cl-5-CF ₃ -2-pyridyl	2-Br-4-CF ₃ -thiazol-5-yl	72-3
51	3-Cl-5-CF ₃ -2-pyridyl	5-Cl-3-benzyloxy-2-benzo[b]furyl	155-7
52	3-Cl-5-CF ₃ -2-pyridyl	3-MeO-2-benzo[b]furyl	oil
53	3-Cl-5-CF ₃ -2-pyridyl	3-Pr ⁱ O-2-benzo[b]furyl	oil
54	3-Cl-5-CF ₃ -2-pyridyl	3-BzO-2-benzo[b]furyl	oil
55	3-Cl-5-CF ₃ -2-pyridyl	3,6-diMeO-2-benzo[b]furyl	oil
56	3-Cl-5-CF ₃ -2-pyridyl	3-BzO-6-MeO-2-benzo[b]furyl	oil
57	3-Cl-5-CF ₃ -2-pyridyl	5-Cl-3-MeO-2-benzo[b]furyl	oil

Cmp	A ¹	A ²	m.p.(°C)
58	3-Cl-5-CF ₃ -2-pyridyl	4-pyridyl	154
59	3-Cl-5-CF ₃ -2-pyridyl	phenylcyclopropyl	oil
60	3-Cl-5-CF ₃ -2-pyridyl	4-morpholinyl	oil
61	3-Cl-5-CF ₃ -2-pyridyl	1-(Bu ^t OC(=O)-pyrrolidin-2-yl	oil

The ¹H N.M.R. data of those compounds in Table A which were not solid at room temperature are presented below.

5 Compound 1

¹H N.M.R. (CDCl₃) δ(ppm) 4.9 (2H, d), 6.55 (1H, m), 7.2 (1H, m), 7.5 (1H, s), 7.8 (1H, br.s), 8.0 (1H, s) and 8.8 (1H, s).

Compound 2

10 ¹H N.M.R. (CDCl₃) δ(ppm) 4.9 (2H, d), 7.1 (1H, s), 7.5 (1H, m), 7.6 (1H, br.s), 7.65 (1H, m), 8.0 (1H, s) and 8.8 (1H, s).

Compound 3

15 ¹H N.M.R. (CDCl₃) δ(ppm) 2.5 (3H, s), 2.7 (3H, s), 4.9 (2H, d), 7.4 (1H, br.s), 8.0 (1H, s) and 8.8 (1H, s).

Compound 4

20 ¹H N.M.R. (CDCl₃) δ(ppm) 3.05 (3H, s), 4.9 (2H, d), 7.7 (1H, br.s), 8.0 (1H, s) and 8.8 (1H, s).

Compound 5

¹H N.M.R. (CDCl₃) δ(ppm) 4.9 (2H, d), 7.45 (1H, d), 7.8 (1H, br.s), 8.0 (1H, s), 8.2 (1H, m), 8.8 (1H, s) and 8.9 (1H, d).

Compound 6

25 ¹H N.M.R. (CDCl₃) δ(ppm) 4.1 (3H, s), 4.9 (2H, d), 7.7 (1H, br.s), 8.0 (1H, s), 8.2 (1H, s), 8.6 (1H, s) and 8.8 (1H, s).

Compound 7

¹H N.M.R. (CDCl₃) δ(ppm) 4.9 (2H, d), 7.7 (1H, br.s), 8.0 (1H, s), 8.7 (1H, s) and 9.0 (1H, s).

Compound 8

5 ¹H N.M.R. (CDCl₃) δ(ppm) 4.95 (2H, d), 7.4-7.65 (5H, m), 8.0 (1H, s), 8.1 (1H, s) and 8.8 (1H, s).

Compound 9

10 ¹H N.M.R. (CDCl₃) δ(ppm) 4.6 (2H, s), 4.9 (2H, d), 6.9-7.1 (3H, m), 7.3-7.4 (2H, m), 7.95 (1H, s), 8.1 (1H, br.s) and 8.8 (1H, s).

Compound 10

15 ¹H N.M.R. (CDCl₃) δ(ppm) 4.95 (2H, d), 7.45 (1H, m), 7.85 (1H, s), 8.0 (1H, s), 8.25 (1H, m), 8.8 (2H, m) and 9.2 (1H, s).

Compound 11

¹H N.M.R. (CDCl₃) δ(ppm) 4.9 (2H, d), 7.0 (1H, d), 7.5 (1H, d), 8.0 (1H, s), 8.6 (1H, s) and 8.8 (1H, s).

20 Compound 12

¹H N.M.R. (CDCl₃) δ(ppm) 5.05 (2H, d), 7.85-8.0 (3H, m), 8.2-8.3 (2H, m), 8.85 (1H, s), 9.2 (1H, s) and 9.7 (1H, d).

Compound 13

25 ¹H N.M.R. (CDCl₃) δ(ppm) 5.0 (2H, d), 7.55 (2H, m), 7.85 (1H, m), 7.95 (1H, m), 8.0 (1H, s) and 8.85 (1H, s)

Compound 14

¹H N.M.R. (CDCl₃) δ(ppm) 2.75 (3H, s), 4.1 (3H, s), 5.0 (2H, d), 8.0 (1H, s), 8.75 (1H, s) and 8.9 (1H, s).

30

Compound 15

¹H N.M.R. (CDCl₃) δ(ppm) 4.50 (2H, d), 7.3 (4H, m), 7.5 (2H, t), 7.95 (1H, s), 8.25 (1H, m), 8.55 (1H, br.s), 8.6 (1H, s) and 8.7 (1H, s).

Compound 16

¹H N.M.R. (CDCl₃) δ(ppm) 2.75 (3H, s), 4.9 (2H, d), 6.8 (1H, s), 7.35 (2H, d), 7.65 (2H, d), 8.0 (1H, s) and 8.8 (1H, s).

5

Compound 32

¹H N.M.R. (CDCl₃) δ(ppm) 1.75 (6H, d, 2xMe), 3.92 (6H, d, 2xMe), 4.89 (2H, d, 2xMe), 7.20-30 (4H, m, Ar), 7.85 (1H, s, NH), 7.95 (1H, s, py-H) and 8.70 (1H, s, py-H).

10

Compound 45

¹H N.M.R. (CDCl₃) δ(ppm) 4.92 (2H, d, -CH₂-), 7.51 (1H, br.s, NH), 7.61 (1H, d, pyH), 8.00 (1H, s, py-H), 8.90 (1H, d, py-H) and 8.96 (1H, s, pyH).

Compound 46

15 ¹H N.M.R. (CDCl₃) δ(ppm) 3.98 (3H, s), 4.82 (2H, d), 6.80 (s, Ar-H), 7.36 (1H, br.s), 7.97 (s, Ar-H) and 8.77 (s, ArH).

Compound 47

20 ¹H N.M.R. (DMSO) δ(ppm) 4.73 (2H, d), 7.00 (s, Ar-H), 8.46 (ArH, s), 8.76 (NH, t), 8.91 (1H, s, Ar) and 12.75 (br.s, NH).

Compound 52

¹H N.M.R. (CDCl₃) δ(ppm) 4.45 (3H, s), 5.00 (2H, d), 7.35 (1H, t), 7.45 (1H, t), 7.60 (1H, d), 7.80 (1H, d), 8.00 (1H, s), 8.40 (1H, t) and 8.80 (1H, s).

25

Compound 53

¹H N.M.R. (CDCl₃) δ(ppm) 1.57 (6H, d), 5.00 (2H, d), 5.15 (1H, m), 7.35 (1H, t), 7.45 (1H, t), 7.60 (1H, d), 7.75 (1H, d), 8.00 (1H, s), 8.70 (1H, t) and 8.80 (1H, s).

Compound 54

30 ¹H N.M.R. (CDCl₃) δ(ppm) 4.95 (2H, d), 5.60 (2H, s), 7.35 (1H, t), 7.40-7.60 (7H, m), 7.80 (1H, d), 7.95 (1H, s), 8.30 (1H, s) and 8.55 (1H, t).

Compound 55

¹H N.M.R. (CDCl₃) δ(ppm) 3.90 (3H, s), 4.35 (3H, s), 5.00 (2H, d), 6.95 (1H, dd), 7.05 (1H, d), 7.65 (1H, d), 8.00 (1H, s), 8.55 (1H, t) and 8.80 (1H, s).

Compound 56

5 ¹H N.M.R. (CDCl₃) δ(ppm) 3.90 (3H, s), 4.95 (2H, d), 5.60 (2H, s), 6.95 (1H, dd), 7.05 (1H, d), 7.40 (3H, m), 7.55 (2H, m), 7.65 (1H, d), 7.95 (1H, s), 8.3 (1H, s) and 8.40 (1H, t).

Compound 57

10 ¹H N.M.R. (CDCl₃) δ(ppm) 4.40 (3H, s), 5.00 (2H, d), 7.40 (1H, d), 7.50 (1H, d), 7.80 (1H, s), 8.00 (1H, s), 8.40 (1H, t) and 8.80 (1H, s).

Compound 59

15 ¹H N.M.R. (CDCl₃) δ(ppm) 1.3 (1H, m), 1.65 (1H, m), 1.85 (1H, m), 2.55 (1H, m), 4.75 (2H, d), 7.1-7.3 (5H, m), 7.95 (1H, s) and 8.7 (1H, s).

Compound 60

20 ¹H N.M.R. (CDCl₃) δ(ppm) 3.4 (4H, dd), 3.8 (4H, dd), 4.75 (2H, d), 6.1 (1H, br.s), 7.95 (1H, s) and 8.95 (1H, s).

Compound 61

¹H N.M.R. (CDCl₃) δ(ppm) 5.05 (2H, d), 7.4-7.6 (4H, m), 7.8 (1H, d), 7.9-8.05 (3H, m), 8.4 (1H, d) and 8.75 (1H, s).

25 Example 3

N-[1-(3-Chloro-5-trifluoromethyl-2-pyridyl)-1-cyanoethyl]-2,6-dichlorobenzamide

Compound 110

30 To a suspension of 1-(3-chloro-5-trifluoromethyl-2-pyridyl)-1-cyanoethylammonium chloride (0.51 g) in dry dichloromethane (10 ml) was added dry triethylamine (0.3 ml) followed by dropwise addition of 2,6-dichlorobenzoyl chloride (0.42 g) and the mixture was stirred for 4 hours. The reaction mixture was washed with aqueous potassium carbonate solution (2x10 ml) and the organic phase was dried (MgSO₄). The filtrate was evaporated onto silica and purified by silica gel chromatography gradient eluting with diethyl ether/dichloromethane (0-20%) to give the title compound, m.p. 166-7 °C.

Preparation of starting materialsa) (3-Chloro-5-trifluoromethyl-2-pyridyl)[(diphenylmethylene)amino]acetonitrile

To a suspension of 60% sodium hydride (4.0 g) in dry dimethylformamide at 0-2 °C under nitrogen was dropwise added a solution of *N*-

(diphenylmethylene)aminoacetonitrile (11.1 g) in dry dimethylformamide (60 ml) and the mixture was stirred for 1 hour at 0 °C. 2,3-Dichloro-5-trifluoromethyl pyridine (7 ml) in dry dimethylformamide (20 ml) was added dropwise over 10 minutes. The mixture was stirred at 0 °C for 30 minutes and then warmed to 22 °C over 3 hours. The mixture was re-cooled to less than 10 °C, and ethanol (3 ml) was added dropwise and stirring continued for 15 minutes.

The mixture was poured as a thin stream into a stirred mixture of diethyl ether (500 ml) and 20% saturated aqueous ammonium chloride solution. The phases were separated and the organic phase was washed with 20% saturated aqueous ammonium chloride solution (2x150 ml). The organic phase was dried over anhydrous magnesium sulphate, filtered and evaporated onto flash silica (50 g). Chromatography over silica eluting with 5-20% diethyl ether in 40/60° Bp petrol gave the title compound, m.p. 108-110 °C.

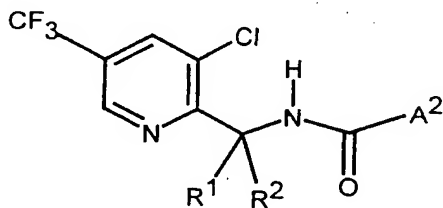
b) 2-(3-Chloro-5-trifluoromethyl-2-pyridyl)-2-[(diphenylmethylene)amino]propionitrile

To a stirred solution of potassium *tert*-butoxide (1.91 g) in dry tetrahydrofuran (50 ml) at -60 °C under nitrogen was dropwise added a solution of the product from stage a) above (5 g) in dry tetrahydrofuran (20 ml). The mixture was stirred at -60 °C for 15 minutes then methyl iodide (1.5 ml) was added dropwise and the mixture warmed to 22°C over 18 hours. The solvent was evaporated *in vacuo* and the residue was partitioned between diethyl ether and 50% saturated aqueous ammonium chloride. The aqueous phase was ether extracted (2x50 ml) and the pooled organic extracts were dried over anhydrous magnesium sulphate. The filtered organic phase was evaporated onto flash silica (20 g). Chromatography over silica eluting with 10-30% diethyl ether in light petroleum (b.p. 40-60 °C) gave the title compound, ¹H N.M.R. CDCl₃ δ (ppm) 2.24 (3H, s, CH₃), 7.10-7.62 (10H, m, Ar-H), 7.90 (1H, s, py-H) and 8.54 (1H, s, py-H).

c) 2-Amino-2-(3-chloro-5-trifluoromethyl-2-pyridyl)propionitrile hydrochloride

To a vigorously stirred solution of the product from stage b) (5.5 g) in diethyl ether (100 ml) under nitrogen was added 2M aqueous hydrogen chloride (100 ml) and stirring continued for 36 hours. The phases were separated and the organic phase was extracted with 2M aqueous hydrogen chloride (2x20 ml). The combined aqueous phases were extracted with diethyl ether (2x20 ml) and the organic extracts discarded. The aqueous phase was evaporated *in vacuo* then azeotroped with toluene (3x50 ml). Trituration with diethyl ether followed by filtration and vacuum drying gave the title compound, m.p. 165-70 °C.

The following compounds of formula Im (see Table B), i.e. compounds of general formula I where A¹ is 3-Cl-5-CF₃-2-pyridyl, L is -C(=O)- and R³ is hydrogen, may be prepared by methods analogous to those of Examples 1, 2 and/or 3.



(Im)

Table B

Cmp	R ¹	R ²	A ²	m.p.(°C)
101	ethoxycarbonyl	H	5-Cl-6-MeO-3-pyridyl	148-50
102	ethoxycarbonyl	H	2-thienyl	oil
103	Me	H	2-furyl	134
104	Me	H	2-thienyl	121
105	piperidiny	H	4-CF ₃ -3-pyridyl	
106	Me	H	4-morpholinyl	139
107	allyl	H	2,6-diCl-phenyl	122
108	cyano	H	2-Cl-6-F-phenyl	114
109	cyano	H	2-Br-6-Cl-phenyl	179
110	cyano	Me	2,6-diCl-phenyl	166
111	cyano	Me	2-Cl-6-F-phenyl	174
112	cyano	Me	2-Br-6-Cl-phenyl	oil
113	cyano	H	2,4-diCl-phenyl	110
114	cyano	Me	2,4-diCl-phenyl	160
115	cyano	H	4-Cl-phenyl	oil
116	cyano	H	3,5-diCl-phenyl	135
117	cyano	Me	3,4-diCl-phenyl	207
118	cyano	Me	3,4-diCl-phenyl	207
119	cyano	Me	4-F-phenyl	201
120	hydroxy	H	4,5-diCl-phenyl	oil

- 5 The ¹H N.M.R. data of those compounds in Table B which were not solid at room temperature are presented below.

Compound 102

¹H N.M.R. (CDCl₃) δ(ppm) 1.23 (1H, t), 4.25 (1H, m), 6.46 (1H, d), 7.12 (1H, m), 7.44
10 (1H, d), 7.50 (1H, d), 7.63 (1H, d) and 8.78 (1H, s).

Compound 112

¹H N.M.R. (CDCl₃) δ(ppm) 2.32 (3H, s, Me), 7.22 (1H, m, Ar-H), 7.40 (1H, d, Ar-H),
15 7.54 (1H, d, Ar-H), 8.14 (1H, d, py-H), 8.36 (1H, s, NHCO) and 8.76 (1H, d, py-H).

Compound 115

¹H N.M.R. (CDCl₃) δ(ppm) 6.54 (1H, d, CHCN), 7.46 (2H, m, 2xAr-H), 7.80 (2H, m, 2xAr-H), 7.9 (1H, d, NHCO), 8.12 (1H, d, py-H) and 8.80 (1H, d, py-H).

5 Compound 120

¹H N.M.R. (CDCl₃) δ(ppm) *inter alia* 7.6 (1H, d, Ar-H), 7.75 (2H, d, 2xAr-H), 8.18 (1H, d, py-H), 8.84 (1H, d, pyH).

Example 410 N-[(3-Chloro-5-trifluoromethyl-2-pyridyl)methyl]-N-[(cyanoimino)methyl]-4-chlorobenzamide (Compound 205)

This compound was prepared in analogous fashion to Example 1 using 2-chlorobenzoyl chloride and the starting material described below.

15 Preparation of Starting materialsN-Cyano-N'-(3-chloro-5-trifluoromethyl-2-pyridylmethyl)formamidine

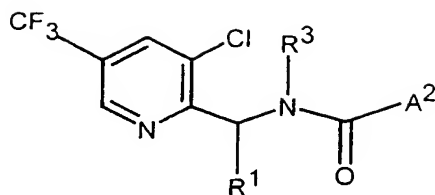
To a stirred suspension of (3-chloro-5-trifluoromethyl-2-pyridyl)methylammonium hydrochloride (19 g) in ethanol (180 ml) was added triethylamine (10.7 ml) and stirring was continued for 15 minutes.

20 Ethoxycyanoimide (8.29 g) in ethanol (20 ml) was then added dropwise and stirred at room temperature for 20 minutes. The solvent was removed *in vacuo* and the residue partitioned between diethyl ether and water. The organic layer was separated and filtered. The filtrate was dried (MgSO₄) and evaporated to give the title product, m.p. 106-8 °C.

25

The following compounds of formula In (see Table C), i.e. compounds of general formula I where A¹ is 3-Cl-5-CF₃-2-pyridyl, L is -C(=O)- and R² is hydrogen, may be prepared by methods analogous to those of Examples 1, 2, 3 and/or 4.

24



(In)

Table C

Cmp	R ¹	R ³	A ²	m.p.(°C)
201	ethoxycarbonyl	Et	2-thienyl	oil
202	methoxycarbonyl	Pr	2-furyl	
203	ethoxycarbonyl	Me	2-thienyl	
204	H	<i>N</i> -cyano-iminomethyl	4-Bu ^l -phenyl	
205	H	<i>N</i> -cyano-iminomethyl	4-Cl-phenyl	oil
206	H	<i>N</i> -cyano-iminomethyl	2-CF ₃ -phenyl	oil
207	H	<i>N</i> -cyano-iminomethyl	4-CF ₃ O-phenyl	
208	H	<i>N</i> -cyano-iminomethyl	4-CF ₃ O-phenyl	
209	H	<i>N</i> -cyano-iminomethyl	4-CF ₃ O-phenyl	isomer of 208
210	H	<i>N</i> -cyano-iminomethyl	3,5-diCl-phenyl	
211	H	<i>N</i> -cyano-iminomethyl	3-Pr ⁱ O-phenyl	
212	H	<i>N</i> -cyano-iminomethyl	3-PhO-phenyl	
213	H	<i>N</i> -cyano-iminomethyl	4-biphenyl	
214	H	<i>N</i> -cyano-iminomethyl	2-tolyl	
215	H	<i>N</i> -cyano-iminomethyl	3-CN-phenyl	

The ¹H N.M.R. data of those compounds in Table B which were not solid at room temperature are presented below.

5

Compound 205

¹H N.M.R. (CDCl₃) δ(ppm) 5.24-5.36 (2H, s, CH₂), 7.45-7.56 (4H, m, Ar-H), 7.82-7.88 (1H, s, Ar-H), 8.50-8.56 (1H, s, N=CH) and 8.84-8.96 (1H, m, ArH).

Compound 206

¹H N.M.R. (CDCl₃) δ(ppm) 5.3-5.5 (2H, m, CH₂), 7.6-7.8 (4H, m, Ar-H), 7.9 (1H, s), 8.5 (1H, m, N=CH) and 8.65 (1H, s, ArH).

5 Example 5N-[(3-Chloro-5-trifluoromethyl-2-pyridyl)methyl]-2-nitrophenylacetamide oxime
(Compound 304)

This compound was prepared in analogous fashion to Example 1 replacing furoyl chloride with 2-nitro-α-chlorobenzaldoxime (see stage b below).

10

Preparation of Starting Materialsa) 2-Nitrobenzaldoxime

To a solution of 2-nitrobenzaldehyde (15.1 g) and hydroxylamine hydrochloride (6.6 g) in ethanol (110 ml) and water (4 ml) was added sodium acetate (13.6 g) and the mixture was stirred at room temperature for 4 hours. The mixture was poured into water (500 ml) and the mixture filtered to give the title product.

15

b) 2-Nitro-α-chlorobenzaldoxime

Through an ice-cold solution of the product from stage a) (10.7 g) in concentrated hydrochloric acid (60 ml) and water (12.3 ml) was bubbled chlorine gas for one hour. The mixture was then stirred at room temperature overnight. The mixture was filtered to give the title product.

20

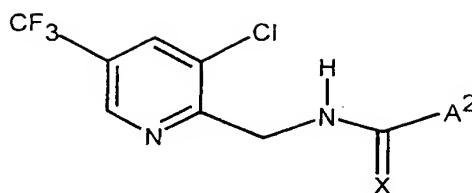
Example 625 N-[(3-Chloro-5-trifluoromethyl-2-pyridyl)methyl]-2-nitrophenylacetamide-O-(phenylcarbamoyl)oxime
(Compound 305)

To a stirred mixture of the product from Example 5 (0.9 g) and 2,6-dichlorophenyl isocyanate (0.33 g) in acetonitrile (50 ml) was added three drops of triethylamine. The mixture was heated to reflux for 2 hours. On cooling the solvent was removed, and the residue purified by silica gel chromatography to give the title product, m.p. 138-40 °C.

30

The following compounds of formula Ip (see Table D), i.e. compounds of general formula I where A¹ is 3-Cl-5-CF₃-2-pyridyl, R¹, R² and R³ are hydrogen and L is
35 -(C=X)-, may be prepared by methods analogous to those of Examples 5 and 6.

26



(Ip)

Table D

Cmp	X	A ²	m.p./°C
301	=NOH	2,6-diCl-phenyl	189-91
302	=NOC(=O)N-Ph	2,6-diCl-phenyl	168-70
303	=NOC(=O)Et	2,6-diCl-phenyl	89-92
304	=NOH	2-NO ₂ -phenyl	122-6
305	=NOC(=O)-2,6-diClPh	2-NO ₂ -phenyl	138-40
306	=NOS(=O) ₂ Me	2-NO ₂ -phenyl	oil

The ¹H N.M.R. data of the compound in Table D which was not solid at room temperature is presented below.

Compound 306

3.1 (3H, s), 4.4 (2H, d), 7.15 (1H, br.t), 7.65 (1H, m), 7.8 (2H, m), 7.8 (2H, m), 7.9 (1H, s), 8.25 (1H, d) and 8.8 (1 H, s).

Example 7

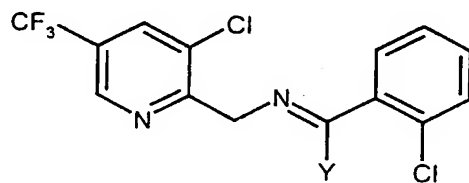
N¹-[(3-Chloro-5-trifluoromethyl-2-pyridyl)methyl]-2-chlorobenzamidine hydrochloride (Compound 307)

This compound was prepared from (3-chloro-5-trifluoromethyl-2-pyridyl)methylammonium hydrochloride and methyl 2-chlorothiobenzimidate hydrogen iodide salt, using methods described in the R C Schnur, *J. Org. Chem.* 1979, Vol.44, No.21, 3726. Methyl 2-chlorothiobenzimidate hydrogen iodide salt was prepared using methods described in Matsuda *et al*, *Synthetic Communications*, 1997, 2393. ¹H N.M.R. (CDCl₃) δ(ppm) 5.10 (2H, d, CH₂), 7.60 (1H, m, ArH), 7.65-7.80 (3H, m, ArH), 8.60 (1H, m, py-H), 9.04 (1H, m, py-H), 9.80-10.00 (2H, br.m, =NH.HCl) and 10.55 (1H, br.m, NH).

Example 8**N-(α -Allylthio-2-chlorobenzylidene)-(3-chloro-5-trifluoromethyl-2-pyridyl)methylamine
(Compound 403)**

To a mixture of sodium hydride (0.33 g) in tetrahydrofuran (10 ml) was added N-[(3-chloro-5-trifluoromethyl-2-pyridyl)methyl]-2-chlorobenzenethioamide (for preparation see PCT/GB/99/00304) (3.07 g) in tetrahydrofuran (50 ml) dropwise with stirring for 20 minutes until effervescence had ceased. Allyl bromide (0.09 g) in tetrahydrofuran (5 ml) was added to the reaction mixture, and the solution was stirred overnight at room temperature. The mixture was evaporated to dryness and the residue partitioned between dichloromethane (10 ml), water (5 ml) and brine (5 ml). The organic phase was separated and the solvent and any residual water evaporated *in vacuo*. The residue was purified by silica gel chromatography, gradient eluting with light petroleum (b.p. 60-80 °C)/diethyl ether to give the title product.

The following compounds of formula IIb (see Table E), i.e. compounds of general formula II where A¹ is 3-Cl-5-CF₃-2-pyridyl, A² is 2-Cl-phenyl and R¹ and R² are hydrogen, may be prepared by methods analogous to those of Example 8.



(IIb)

Table E

Cmp	Y	mass spectral data [m/z (API)]
401	SMe	379 (M+H) ⁺
402	SEt	393 (M+H) ⁺
403	allylthio	405 (M+H) ⁺
404	benzylthio	455 (M+H) ⁺
405	2-Me-benzylthio	469 (M+H) ⁺
406	4-Cl-benzylthio	489 (M+H) ⁺

Example 9

N-{[3-Chloro-5-(trifluoromethyl)-2-pyridyl]methyl}-4,5-dichloro-3,6-epoxy-1,2-dicarboximide

5 (Compound 503)

To a mixture of (3-chloro-5-trifluoromethyl-2-pyridyl)methylamine (1 mmol) in xylene (5 ml) was added triethylamine (1 mmol) at room temperature and the mixture was stirred at room temperature for 1.5 hour. The mixture was filtered and the filtrate was added to 4,5-dichloro-3,6-epoxy-1,2-dicarboxylic anhydride (1 mmol) in xylene (5 ml) at
10 room temperature. The reaction mixture was heated at 130 °C for 48 hours. On cooling, the solvent was removed, the residue was washed with diethyl ether/ light petroleum (b.p. 60-80 °C) and dried to give the title product.

Example 10

15 3-(3-Chloro-5-trifluoromethyl-2-pyridyl)methyl-1,2,3,4-tetrahydro-4-oxo-2-thioxoquinazoline

(Compound 504)

To a suspension of [3-chloro-5-(trifluoromethyl)-2-pyridyl]methylammonium chloride (0.12 g) and 2-(methoxycarbonyl)phenylthioisocyanate (0.10 g) in dry tetrahydrofuran
20 (10 ml) was added 10 drops of triethylamine. The mixture was stirred at room temperature overnight. The solvent was removed by evaporation *in vacuo* and the product was extracted with ethyl acetate and washed with 2M hydrochloric acid. The organic layer was collected and evaporated *in vacuo* to give the title product. ¹H N.M.R. (CDCl₃) δ(ppm) 6.15 (2H, s), 7.15 (1H, d), 7.35 (1H, t), 7.7 (1H, t), 7.9 (1H, s), 8.18
25 (1H, d), 8.75 (1H, s) and 10.2 (1H, s, NH).

Example 11

3-(3-Chloro-5-trifluoromethyl-2-pyridyl)methyl-3,4-dihydro-4-oxo-2-[(4-chlorobenzyl)thio]quinazoline

30 (Compound 507)

This compound was prepared from the product of Example 10 and 4-chlorobenzylbromide in analogous fashion to Example 8. m.p. 137 °C.

Example 12

35 N-[3-Chloro-5-trifluoromethyl-2-pyridyl)methyl]phthalimide

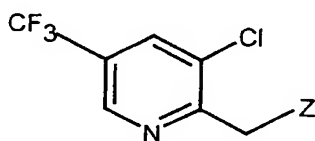
(Compound 518)

A mixture of phthalic anhydride (0.601 g), 3-chloro-5-trifluoromethyl-2-pyridyl)methylammonium hydrochloride (1.0 g) and powdered potassium carbonate (0.28 g) in dimethylformamide was stirred at 148 °C for 7 hours. On cooling, water (10 ml) was added and the mixture filtered to give a solid. The solid was dissolved in ethyl acetate, dried (MgSO₄) and the solvent removed. The residue was triturated from diethyl ether to give the title product, m.p. 145-6 °C.

Example 132-([3-Chloro-5-(trifluoromethyl)-2-pyridyl]methyl)-3-hydroxy-1-indanoneCompound 516

To an ice-cooled solution of the product from Example 12 (1.23 g) in methanol (12.3 ml) was added sodium borohydride portionwise over 5 minutes and stirring was continued overnight. The mixture was partitioned between saturated ammonium chloride (50 ml) and ethyl acetate (50 ml). The layers were separated and the organic layer was dried (MgSO₄). Evaporation gave the title compound, m.p. 174-8 °C.

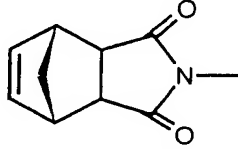
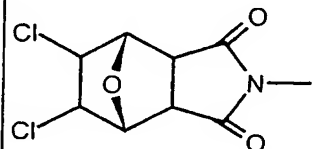
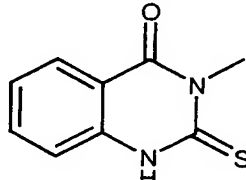
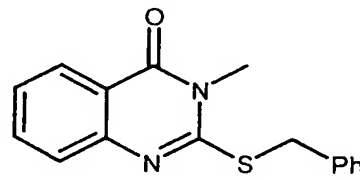
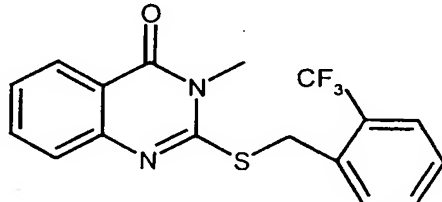
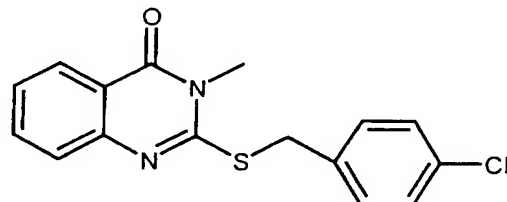
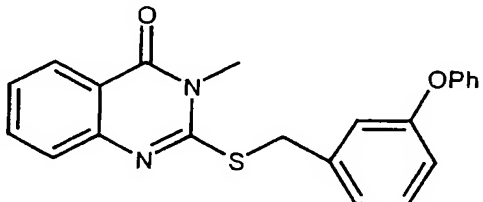
The following compounds of formula Iq (see Table F), i.e. compounds of general formula I where A¹ is 3-Cl-5-CF₃-2-pyridyl, R¹ and R² are hydrogen, and R³ and A² together with the interconnecting atoms form a ring, may be prepared by methods analogous to those of Examples 9, 10, 11, 12 and/or 13. Compounds 514, 515 and 517 were prepared by simple alkylation of compound 516; such methods are familiar to skilled chemists.

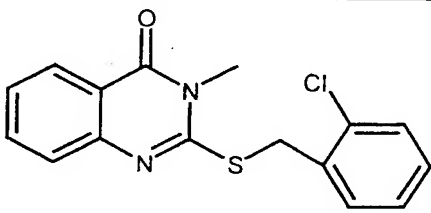
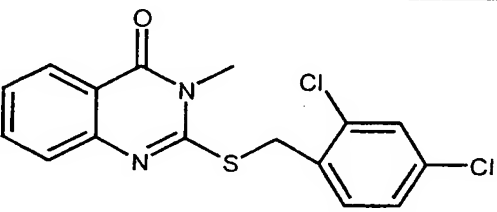
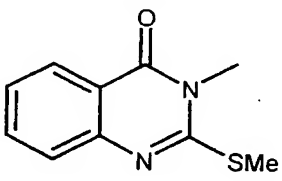
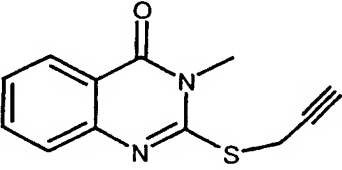
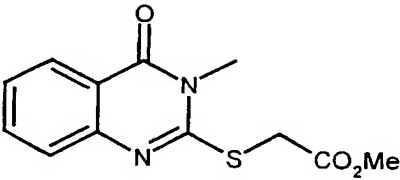
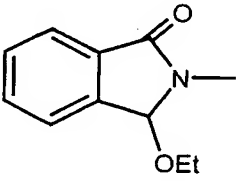
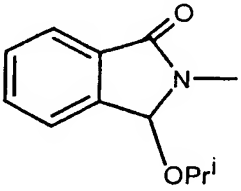
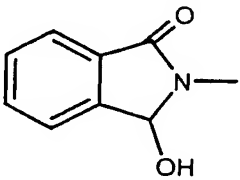


(Iq)

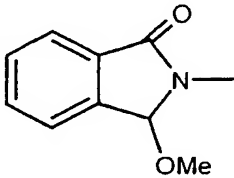
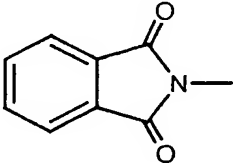
Table F

Cmp	Z	m.p.(°C)
501		126

Cmp	Z	m.p.(°C)
502		oil
503		oil
504		oil
505		148-50
506		134-6
507		137
508		105

Cmp	Z	m.p.(°C)
509		149
510		85-8
511		oil
512		oil
513		oil
514		oil
515		oil
516		174-8

32

Cmp	Z	m.p.(°C)
517		oil
518		145-6

The ^1H N.M.R. data of those compounds in Table D that were not solid at room temperature are presented below.

5 Compound 502

^1H N.M.R. (CDCl_3) δ (ppm) 1.6 (1H, d), 1.8 (1H, d), 2.4 (4H, s), 4.9 (2H, s), 6.15 (2H, s), 7.9 (1H, s) and 8.6 (1H, s).

Compound 503

10 ^1H N.M.R. (CDCl_3) δ (ppm) 3.25 (1H, d), 3.85 (1H, d), 3.95 (1H, d), 4.35 (1H, m), 4.9 (1H, s), 4.95 (2H, s), 5.1 (1H, m), 7.9 (1H, s) and 8.65 (1H, s).

Compound 504

^1H N.M.R. (CDCl_3) δ (ppm) 6.15 (2H, s), 7.15 (1H, d), 7.35 (1H, t), 7.7 (1H, t), 7.9 (1H, s), 8.18 (1H, d), 8.75 (1H, s) and 10.2 (1H, s, NH).

15

Compound 511

^1H N.M.R. (CDCl_3) δ (ppm) 2.7 (3H, s), 5.7 (2H, s), 7.25 (1H, t), 7.65 (1H, d), 7.75 (1H, t), 7.95 (1H, s), 8.25 (1H, d) and 8.6 (1H, s).

20

Compound 512

^1H N.M.R. (CDCl_3) δ (ppm) 2.25 (1H, t), 4.1 (2H, d), 5.65 (2H, s), 7.45 (1H, t), 7.65 (1H, d), 7.75 (1H, t), 7.95 (1H, s), 8.25 (1H, d) and 8.6 (1H, s).

Compound 513

¹H N.M.R. (CDCl₃) δ(ppm) 3.8 (3H, s), 4.05 (2H, s), 5.65 (2H, s), 7.45 (1H, t), 7.55 (1H, d), 7.75 (1H, t), 7.95 (1H, s), 8.25 (1H, d) and 8.6 (1H, s).

Compound 514

¹H N.M.R. (CDCl₃) δ(ppm) 1.1 (3H, t), 3.1 (1H, m), 3.3 (2H, s), 4.6 (1H, d), 5.4 (1H, d), 6.1 (1H, s), 7.4-7.6 (3H, m), 7.9 (2H, m) and 8.6 (1H, s).

Compound 515

¹H N.M.R. (CDCl₃) δ(ppm) 1.1 (3H, d), 1.2 (3H, d), 3.7 (1H, m), 4.7 (1H, d), 5.4 (1H, d), 6.1 (1H, s), 7.4-7.6 (3H, m), 8.8-8.9 (2H, m) and 8.6 (1H, s).

Compound 517

¹H N.M.R. (CDCl₃) δ(ppm) 3.0 (3H, m), 4.6 (1H, d), 5.4 (1H, d), 6.1 (1H, s), 7.5-7.7 (3H, m), 7.9 (2H, m) and 8.6 (1H, s).

Test Example

Compounds were assessed for activity against one or more of the following:

Phytophthora infestans: late tomato blight

Plasmopara viticola: vine downy mildew

Erysiphe graminis f. sp. tritici: wheat powdery mildew

Pyricularia oryzae: rice blast

Leptosphaeria nodorum: glume blotch

Aqueous solutions or dispersions of the compounds at the desired concentration, including a wetting agent, were applied by spray or by drenching the stem base of the test plants, as appropriate. After a given time, plants or plant parts were inoculated with appropriate test pathogens before or after application of the compounds as appropriate, and kept under controlled environmental conditions suitable for maintaining plant growth and development of the disease. After an appropriate time, the degree of infection of the affected part of the plant was visually estimated. Compounds are assessed on a score of 1 to 3 where 1 is little or no control, 2 is moderate control and 3 is good to total

control. At a concentration of 500 ppm (w/v) or less, the following compounds scored 2 or more against the fungi specified.

Phytophthora infestans

5 7, 14, 34, 37, 45, 104, 108, 110, 111, 112, 206, 210 and 214.

Plasmopara viticola:

7, 14, 34, 37, 41, 42, 43, 44, 45, 109, 110, 111, 112, 206, 214 and 301.

10 *Erysiphe graminis f. sp. tritici*:

2, 4, 15, 31, 108 and 516

Pyricularia oryzae

4, 39, 41, 108, 109, 113, 116, 201, 215, 512 and 516.

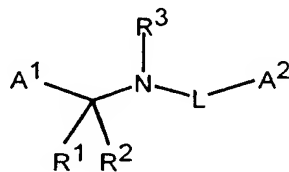
15

Leptosphaeria nodorum

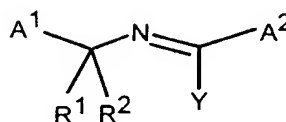
11, 15, 46, 48, 49 and 50.

Claims

- 5 1. The use of compounds of general formula I or II or salts thereof as
phytopathogenic fungicides



(I)



(II)

wherein

A^1 is 2-pyridyl or its *N*-oxide, each of which may be substituted by up to four
10 groups at least one of which is haloalkyl;

A^2 is heterocyclyl or carbocyclyl, each of which may be substituted;

R^1 and R^2 , which may be the same or different, are R^b , cyano, nitro, halogen,
-OR^b, -SR^b or optionally substituted amino, or R^1 and R^2 together with
the carbon to which they are attached may form a 3-, 4-, 5- or 6- carbo-
15 or heterocyclic ring, which may be substituted;

R^3 is R^b , -OR^b, or -N(R^b)₂, cyano, *N*-substituted iminomethyl or nitro; or R^3
and A^2 , together with the interconnecting atoms, may form a 5- or 6-
membered ring;

L is -C(=X)- or -SO₂-, where X is oxygen, sulfur, N-OR^b, N-R^b or N-N(R^b)₂;

and

Y is halogen, -OR^b, -SR^b, -N(R^b)₂, -NR^b(OR^b) or -NR^bN(R^b)₂;

and R^b , which may be the same or different, is alkyl, alkenyl, alkynyl,
carbocyclyl or heterocyclyl, each of which may be substituted; or
hydrogen or acyl, or two adjacent R^b groups together with the
interconnecting atoms, may form a 5- or 6-membered ring;

25 with the proviso that when A^1 is 2-pyridyl, R^1 is hydrogen, R^2 is hydrogen,
optionally substituted alkyl or acyl, L is -C(=X)-

or -SO₂-, X is oxygen or sulfur and R³ is hydrogen or optionally substituted alkyl, A² is not optionally substituted phenyl.

2. A pesticidal composition comprising at least one compound as claimed in claim
5 1 in admixture with an agriculturally acceptable diluent or carrier.
3. A method of combating pests at a locus infested or liable to be infested therewith, which comprises applying to the locus a compound as claimed in claim 1.

INTERNATIONAL SEARCH REPORT

International Application No
PCT/EP 00/08269

A. CLASSIFICATION OF SUBJECT MATTER

IPC 7 A01N43/40 A01N43/82 A01N43/80 A01N43/54 A01N43/56
A01N43/90 A01N43/50 A01N43/78 A01N43/42 A01N43/60
A01N53/00 A01N47/38 A01N47/40 A01N47/24

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

IPC 7 A01N

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practical, search terms used)

WPI Data, CHEM ABS Data, EPO-Internal

C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	EP 0 329 020 A (MITSUBISHI CHEM IND) 23 August 1989 (1989-08-23) page 4, line 23 - page 5, line 15 page 5, line 45 page 5, line 52 page 8, line 19 - line 21	1-3
Y	---	1-3
Y	WO 97 08135 A (BAYER AG ; SEITZ THOMAS (DE); NAUMANN KLAUS (DE); TIEMANN RALF (DE)) 6 March 1997 (1997-03-06) page 1, line 16 - line 25 page 2, line 9 - line 17 page 6, line 4 - line 6 page 6, line 13 - line 16 page 8, line 21 page 8, line 29 ---	1-3
	--- -/-	

☒ Further documents are listed in the continuation of box C.

☒ Patent family members are listed in annex.

* Special categories of cited documents:

A document defining the general state of the art which is not considered to be of particular relevance

E earlier document but published on or after the international filing date

L document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)

O document referring to an oral disclosure, use, exhibition or other means

P document published prior to the international filing date but later than the priority date claimed

T later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention

X document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone

Y document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art.

& document member of the same patent family

Date of the actual completion of the international search

6 December 2000

Date of mailing of the international search report

21/12/2000

Name and mailing address of the ISA

European Patent Office, P.B. 5818 Patentlaan 2
NL - 2280 HV Rijswijk
Tel. (+31-70) 340-2040, Tx. 31 651 epo nl,
Fax: (+31-70) 340-3016

Authorized officer

Lamers, W

INTERNATIONAL SEARCH REPORT

Inter. Application No
PCT/EP 00/08269

C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
Y	US 3 609 158 A (TORBA FLORENCE E) 28 September 1971 (1971-09-28) column 1, line 12 - line 66 ----	1-3
Y	WO 99 07687 A (AGREVO UK LTD ; COOPER IAN PAUL (GB); WEST PETER JOHN (GB); CARVER) 18 February 1999 (1999-02-18) page 1, line 1 - page 5, line 29 page 28, compound 53b ----	1-3
Y	EP 0 726 266 A (MITSUBISHI CHEM CORP) 14 August 1996 (1996-08-14) page 2, line 48 - page 3, line 15 page 3, line 54 page 4, line 54 - line 56 ----	1-3
Y	US 4 423 222 A (ASH MARY L ET AL) 27 December 1983 (1983-12-27) column 1, line 6 - line 25 ----	1-3
Y	EP 0 334 138 A (BAYER AG) 27 September 1989 (1989-09-27) page 2, line 1 - line 28 page 3, line 33 page 4, line 1 - line 2 page 4, line 40 page 9, line 6 - line 8 page 14, line 1 - line 31 ----	1-3
Y	EP 0 404 190 A (TAKEDA CHEMICAL INDUSTRIES LTD) 27 December 1990 (1990-12-27) page 2, line 38 - page 4, line 53 page 13, line 5 - line 7 page 13, line 26 - line 35 page 17, line 31 - line 40 ----	1-3
A	DATABASE CHEMABS 'Online! CHEMICAL ABSTRACTS SERVICE, COLUMBUS, OHIO, US; retrieved from STN-INTERNATIONAL, accession no. 123:340114 CA XP002154811 abstract IT: 170788-47-3 & JP 07 173139 A (MITSUBISHI) 11 July 1995 (1995-07-11) ----- -/--	1-3

INTERNATIONAL SEARCH REPORT

International Application No

PCT/EP 00/08269

C.(Continuation) DOCUMENTS CONSIDERED TO BE RELEVANT		
Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
A	<p>DATABASE CHEMABS 'Online! CHEMICAL ABSTRACTS SERVICE, COLUMBUS, OHIO, US; retrieved from STN-INTERNATIONAL, accession no. 125:275659 CA XP002154812 abstract IT: 182225-13-4 & JP 08 208615 A (DAINIPPON) 13 August 1996 (1996-08-13)</p>	1-3
A	<p>DATABASE CHEMABS 'Online! CHEMICAL ABSTRACTS SERVICE, COLUMBUS, OHIO, US; retrieved from STN-INTERNATIONAL, accession no. 111:77653 CA XP002154813 abstract IT: 121905-96-2, 121905-95-1 & JP 64 003162 A (TOKUYAMA SODA) 6 January 1989 (1989-01-06)</p>	1-3
P,A	<p>WO 99 42447 A (MOLONEY BRIAN ANTHONY ;SAVILLE STONES ELIZABETH ANNE (GB); AGREVO) 26 August 1999 (1999-08-26) page 1, line 3 - line 28</p>	1-3

INTERNATIONAL SEARCH REPORT

information on patent family members

International Application No

PCT/EP 00/08269

Patent document cited in search report		Publication date	Patent family member(s)	Publication date
EP 0329020	A	23-08-1989	BR 8900743 A DE 68900029 D JP 2062876 A JP 2751309 B US 4968805 A	17-10-1989 07-03-1991 02-03-1990 18-05-1998 06-11-1990
WO 9708135	A	06-03-1997	DE 19626311 A AU 6874096 A BR 9610048 A CN 1200725 A CZ 9800610 A EP 0848700 A HU 9802859 A JP 11511442 T PL 325277 A SK 27198 A US 6001879 A	23-10-1997 19-03-1997 06-07-1999 02-12-1998 17-06-1998 24-06-1998 29-03-1999 05-10-1999 20-07-1998 09-09-1998 14-12-1999
US 3609158	A	28-09-1971	US 3705170 A US 3711486 A US 3787420 A	05-12-1972 16-01-1973 22-01-1974
WO 9907687	A	18-02-1999	AU 8635298 A	01-03-1999
EP 0726266	A	14-08-1996	JP 8277287 A AT 161841 T AU 4331196 A CN 1137523 A DE 69600134 D DE 69600134 T ES 2112656 T US 5705453 A	22-10-1996 15-01-1998 15-08-1996 11-12-1996 12-02-1998 20-05-1998 01-04-1998 06-01-1998
US 4423222	A	27-12-1983	NONE	
EP 0334138	A	27-09-1989	DE 3835660 A BR 8901384 A JP 1305074 A US 4959379 A	12-10-1989 07-11-1989 08-12-1989 25-09-1990
EP 0404190	A	27-12-1990	AT 129498 T AU 622330 B AU 5777490 A BR 9002976 A CA 2019651 A DE 69023171 D DE 69023171 T HU 54685 A PL 165356 B PL 164890 B US 5498774 A JP 4018091 A	15-11-1995 02-04-1992 17-01-1991 20-08-1991 23-12-1990 30-11-1995 28-03-1996 28-03-1991 30-12-1994 31-10-1994 12-03-1996 22-01-1992
JP 07173139	A	11-07-1995	NONE	
JP 08208615	A	13-08-1996	NONE	
JP 64003162	A	06-01-1989	NONE	

INTERNATIONAL SEARCH REPORT

information on patent family members

International Application No

PCT/EP 00/08269

Patent document cited in search report	Publication date	Patent family member(s)	Publication date
WO 9942447 A	26-08-1999	AU 2527199 A	06-09-1999
		EP 1056723 A	06-12-2000
		NO 20004159 A	17-10-2000
<hr/>			